Biosciences for for an Expanding Bioeconomy

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Biosciences at a Glance

3 Scientific Divisions and a User Facility

- Biological Systems and Engineering
- Environmental Genomics and Systems Biology
- Molecular Biophysics and Integrated Bioimaging
- Joint Genome Institute

670+ staff and 733 affiliates



KBase

212

147

66

68

83

29

67

733

\$180+M in funding in FY2018





Scientific

Technical

Faculty

PostDoc

Graduate

Affiliates

Undergraduate

Operations





Berkeley Lab's Strategic Biosciences Area

10-Year Biosciences Strategic Plan released in 2013

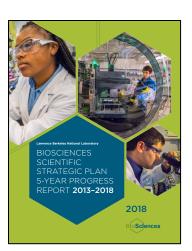
- Energy
- Environment
- Health
- Biomanufacturing
- Technologies

BSP is structured for implementation

- Progress can be measured
 - -Five 10-year goals
 - -10-year metrics of success
 - -5-year milestones

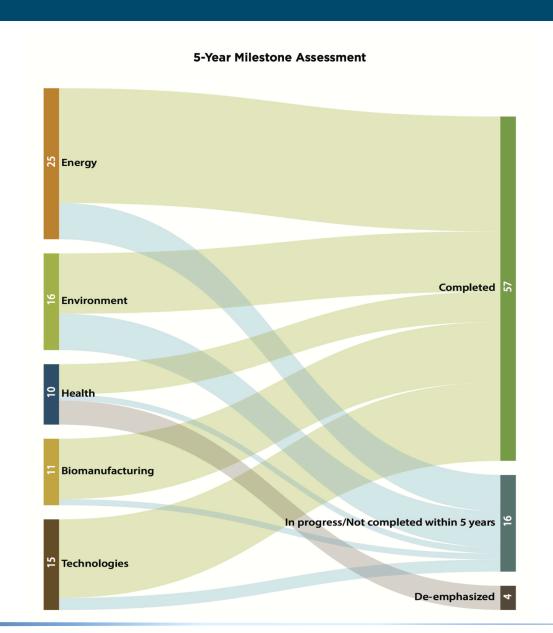
Progress report completed in May 2018







Biosciences Completed 74% of Our Milestones

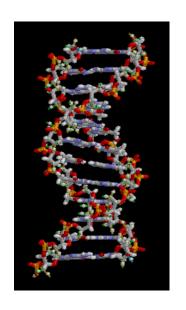


Pillars of the U.S. Bioeconomy

BIOMASS

BIOTECHNOLOGY





"BIO" = DNA



「CTCGTCCGTAACGATCTGGGGGGCAAAACCGAATATCCGTATTCTCGTCCTACGGGTCCACAATGAGAAAGTCO 「GCGCGTGATCGTCAGTTAAGTTAAATTAATTCAGGCTACGGTAAACTTGTAGTGAGCTAAGAATCACGGGAATC \CGGGTTCGCTACAGATGAACTGAATTTATACACGGACAACTCATCGCCCATTTGGGCGTGGGCACCGCAGATCA \AAGTGGCAGATTAGGAGTGCTTGATCAGGTTAGCAGGTGGACTGTATCCAACAGCGCATCAAACTTCAATAAAT CAAAGCGTTGTAGTGGTCTAAGCACCCCT !ATCGTTAGCGTAGTACAACCCTTCCCCCTTG ACGTTCAATAAGAGGGGCTCTACAGCGCCGC \GGTGCGACATGGGGCCAGTTAGCCTGCC0 FTTTTAAATTAGGATGCCGACCCCATCATT ICATAGATATTTCTTCAGGAGTAATAGCGACA \GCTGACACGCAAGGGTCAACAATAATTT0 BTGAACGACTGTCTTTGCAAGAACCAACTGGG CTTAGATTCGCGTCCTAACGTAGTGAGGG0 IGATCAGGCATGAGAAACCGACGTCGAGTCTA BAAGGATCTCCTACATCAAAGACTACTGGGCG CACACGAGTTGTAAACAACTTGATTGCTAT \TCTGGATCCGAGTCAGAAATACGAGTTAA IGACCGGTGAAAACACGTGCCATGGGTTGCGT \GACCGTAGTCAGAAGTGTGGCGCGCTATT 'GGAGTATACAGAATTGCTCTTCTACGACGTA \GGAGCTCGGTCCCCAATGCACGCCAAAAA 'CAAACTGCGCATGGTCCCTCCGCCGGTGGCA CTATTATCCATCCGAACGTTGAACCTACTT 'GTCCTCAACAGTATCGCTTATGAATCGCATG CGGCTGTGGATCTTAACGGCCACATTCTTA LCCGATCGCCTTTCCTCGCTGGTACAATGAGT ACTAAGTTATCCAGATCAAGGTTTGAACGG TGTGACTGAACCCGGGAGGAAATGCAGAGAA CTGTTTCAAGGCCTCTGCTTTGGTATCACT CAGACAAGTGGCAAAATTTCGTGCGCCTCTO CTAGGTATTCACGCAACCGTCGTAACATGC |GCGCCAGGGGGGCATACTAGGTCCCGGAGCT JohanoGGTTACGACACAATTATCGGGATCGTCTAGA AAAGACTACCCTATGGATTCCTTGGAGCGGGGACAA STGTTGGGTCGGGCAAGTCCCCGAAGCTCGGCCAAAAGATTCGCCATGGAACCGTCTGGTCCTGTTAGCGTGTAO GCCTGCTCCTGTTCCGGGTACCATAGATAGACTGAGATTGCGTCAAAAAATTGCGGCGAAAATAGAGGGGCTCCT [GTAGAAATACCAGACTGGGGAATTTAAGCGCTTTCCACTATCTGAGCGACTAAACATCAACAAATGCGTCTACT CA ATCCGC AGT AGGC AATT AC AACCTGGTTC AGATC ACTGGTT AATC AGGG ATGTCTTC AT AAGATT AT ACTTG

SACAGTAGAGAGCTATTGTGTAATTCAGGCTCAGCATTCATCGACCTTTCCTGTTGTGAATATTGTGCTAATGCA

"Reading" and "Writing" DNA

"Reading" DNA

Human Genome Project

1990-2000

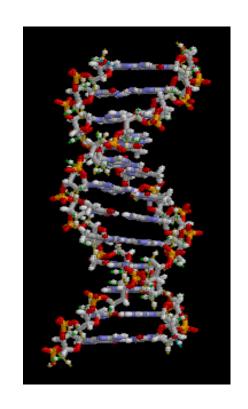
"Writing" DNA

No federal initiative

Today, \$1 buys:

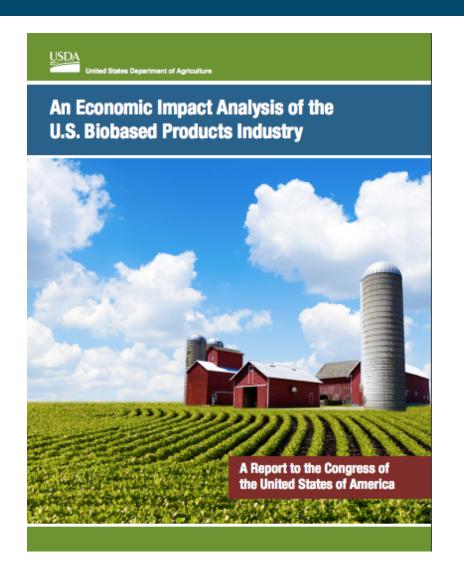
Reading DNA = 500,000 "letters"

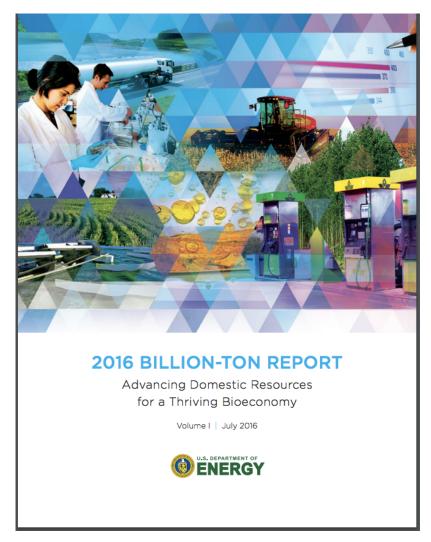
Writing DNA =10 "letters"





U.S. Bioeconomy-related Assessments (non-biomedical)







Policy for the U.S. Bioeconomy 2012



NATIONAL BIOECONOMY BLUEPRINT

April 2012



2016 >\$350B revenues from GMOs >10% annual growth



Support R&D investments that will provide the foundation for the future bioeconomy.



Facilitate the transition of bioinventions from research lab to market, including an increased focus on translational and regulatory sciences.

Develop and reform regulations to reduce barriers, increase the speed and predictability of regulatory processes, and reduce costs while protecting human and environmental health.

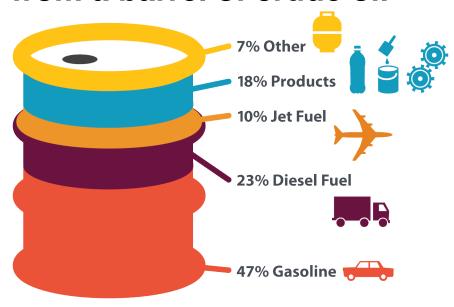
Update training programs and align academic institution incentives with student training for national workforce needs.

Identify and support opportunities for the development of public-private partnerships and precompetitive collaborations—where competitors pool resources, knowledge, and expertise to learn from successes and failures.

In the U.S. Petroleum Is the Primary Source for Transportation Fuels and Chemicals

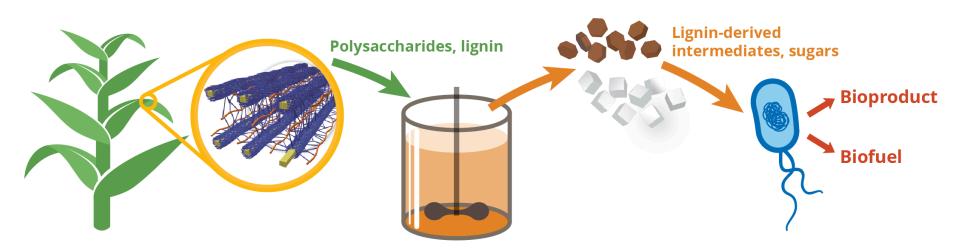


Petroleum products made from a barrel of crude oil



Source: U.S. Department of Energy

Biomass + Biotechnology: engineering of plants and microbes drives the U.S. bioeconomy



Create crops with desired characteristics:

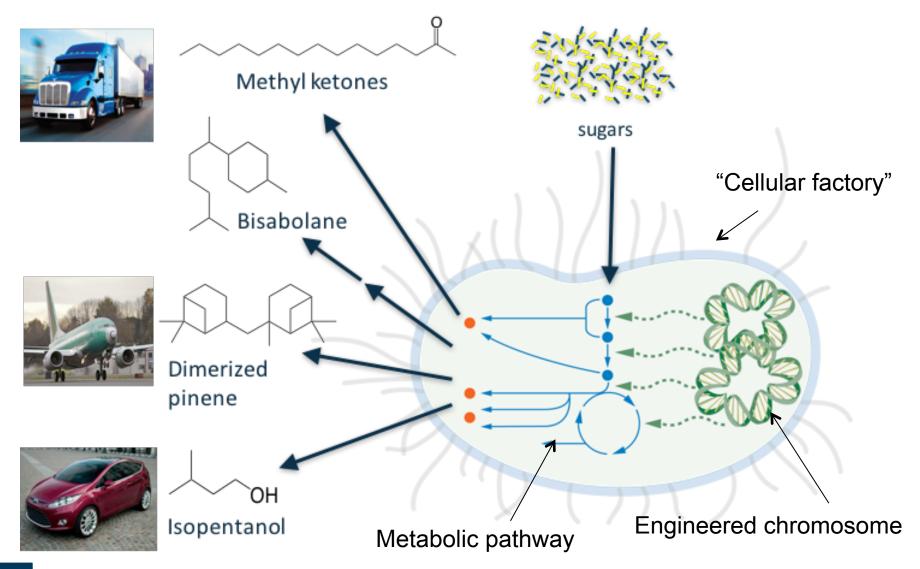
- Increased sugar
- Decreased lignin
- Drought tolerance
- Reduced dependence on fertilizers

Create microbial production hosts:

- Genetic pathways for desired products
- Designed features for fermenter performance

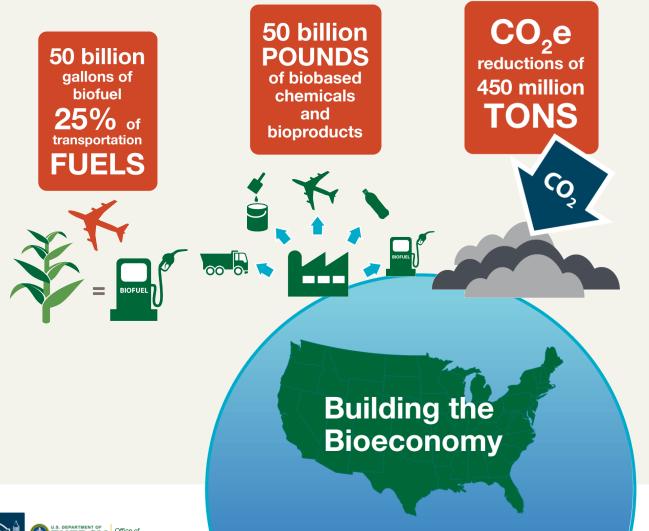


Example: engineered organisms for biofuels





A billion tons of biomass has the potential to produce...



1.1 million
DIRECT
JOBS
and keeps
\$250 billion
in the U.S.

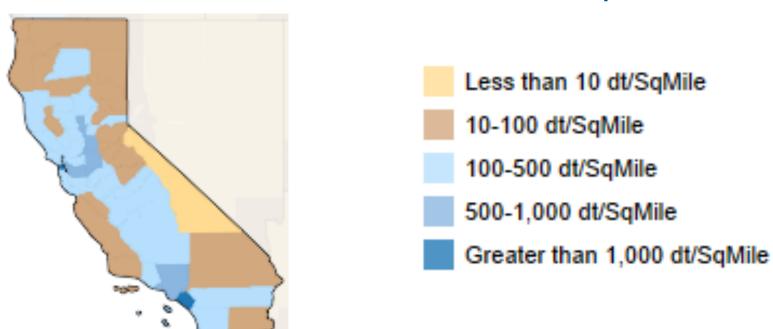


Projections based on:

Rogers, J.N., Stokes, B., Dunn, J., Cai, H., Haq, Z., and Baumes, H. (2016), An Assessment of the Potential Products and Economic and Environmental Impacts Resulting from a Billion Ton Bioeconomy. *Biofuels, BioProd. Bioref.* Doi:10.1002/bbb. 1728

California Bioeconomy: Biomass data

2017 data from the Billion Ton report



Only 15% of available biomass is used!



California "Gold": Diverse biomass resources

- "F.O.G.": fats, oils, greases
- > Small, localized sources: breweries, etc.
- Municipal solid waste
- Animal waste from livestock ranches
- Specialty crop agricultural waste
- Forest waste

The challenge: engineering biology remains slow



The challenge: cost and time to market

Molecule	Company	Cost	Time
1,3-Propanediol (PDO)	DuPont - Tate & Lyle	>\$120M	15 years
Artemisinin	UC Berkeley, Amyris, Sanofi	>\$50M	10 years

Possible savings of *billions* of dollars by reducing development time of products, reducing energy intensity and increasing carbon efficiency



Industry needs: de-risk technologies and decrease time to market



ABOUT

CAPABILITIES

PEOPLE

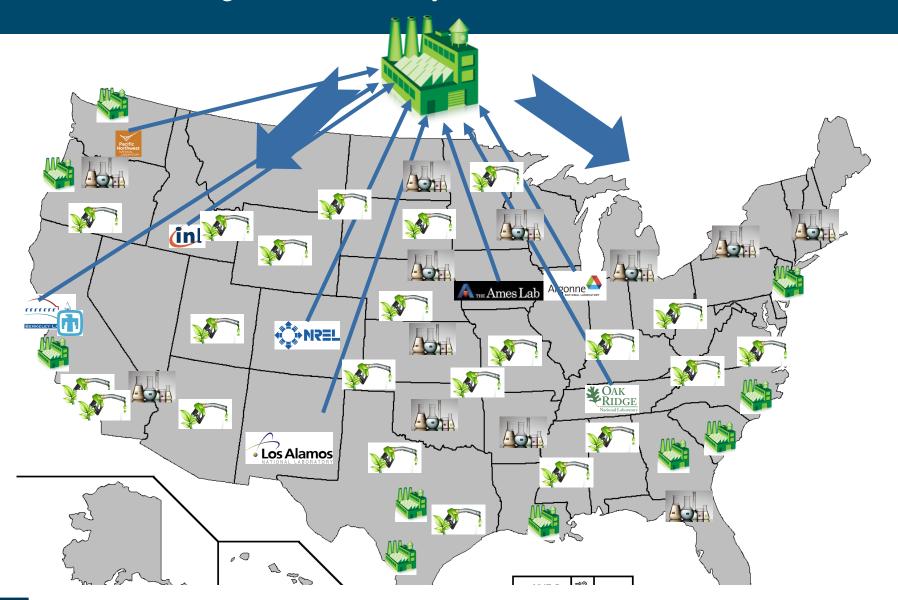
WORK WITH US



- Established in Fiscal Year 2016, a consortium of 8 national laboratories
- Aimed to address shared industry research needs:
 - Accelerate engineering biology and process improvements for biomanufacturing using automation and machine learning
 - Develop and make publicly available standards, tools, technologies
- Democratize "writing DNA"
- \$20M in Fiscal Year 2019 budget; total of \$64.5M



A Distributed Agile BioFoundry: 8 National Laboratories



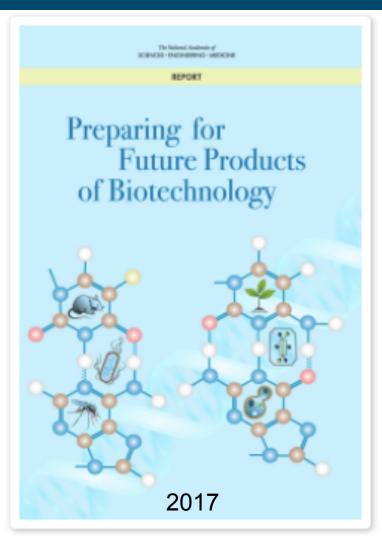


Regulation of Biotechnology

1986 Coordinated Framework for the Regulation of Biotechnology

Updated in 2017

Regulation of products of biotechnology



Biotechnology Regulatory Authorities:

- US Department of Agriculture: engineered plants/crops (Plant Protection Act)
- Food & Drug Administration: engineered animals (Food, Drug & Cosmetic Act)
- Environmental Protection Agency: engineered micro-organisms (Toxic Substances Act)

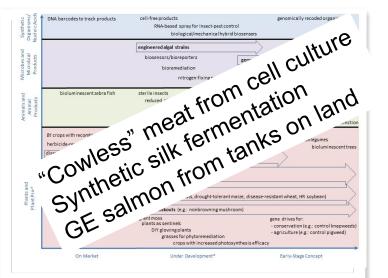
Commissioned an independent study by the U.S. National Academies for horizon-scanning to prepare for new products of biotechnology

What is a Biotechnology Product?

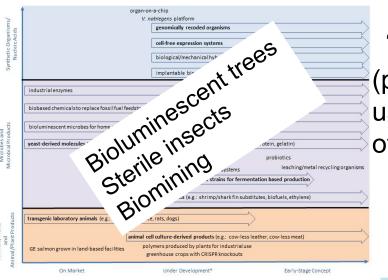
Products developed through genetic engineering or genome engineering or the targeted or *in vitro* manipulation of genetic information of organisms, including plants, animals, and microbes

Includes products where the engineered DNA molecule is itself the "product" as in an engineered molecule used as a DNA information-storage medium

Contained use products



Open release products



"Platforms"
(products
used to make
other products)

The National Academies of





New products will be unfamiliar and complex

A

Domesticated organisms

Transgenic / recombinant DNA

One or only a few genepathway engineering Ample comparators В

Domesticated & Undomesticated organisms

Transgenic, new genome engineering

Multiple pathway engineering

Few to no comparators

C

Many candidate organisms

Genome engineering, gene drives

Genome refactoring, recoding, cellfree synthesis

Few to no comparators

D

Synthetic communities of microbes and synthetic, multicellular plants & animals

Metagenome & microbiome engineering

Population and ecosystem engineering

No or ambiguous comparators

Product Complexity and Novelty

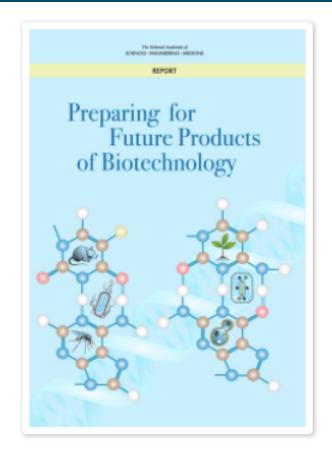


Emergent Themes of the Study

- The bioeconomy is growing rapidly and the U.S. regulatory system needs to provide a balanced approach for consideration of the many stakeholders and competing interests in the face of this expansion
- The profusion of biotechnology products over the next 5-10 years has the potential to overwhelm the U.S. regulatory system
- Regulators will face difficult challenges that go beyond contained industrial uses and traditional environmental release, including new "players"
- The safe use of new biotechnology products requires rigorous, predictable, and transparent risk-analysis processes that mirror the scope, scale, complexity, and tempo of biotechnology development



So...regulatory systems need to be adaptable!



www.nap.edu/catalog/24605/preparing-for-future-products-of-biotechnology

The National Academies of



Berkeley Lab-wide Strategic Priorities 2018







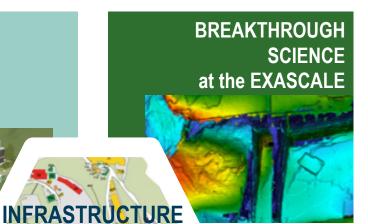






BIOLOGICAL and **ENVIRONMENTAL SCIENCE** at the **BIOCAMPUS**



























Berkeley Lab Infrastructure Priorities Future BioCampus

JBEI+ (envisioned)

- Would support development of sustainable biofuels and bioproducts
- Would house the Joint BioEnergy Institute, the Advanced Biofuels and Bioproducts Process
 Development Unit, and the Agile BioFoundry

Biological and Environmental Program Integration Center (BioEPIC)

- CD-0
- Would bring a wide range of biological and environmental expertise together to focus on soil-microbe-plant interactions



- Currently under construction expected completion 2019
- Will house the Joint Genome Institute and the DOE Systems Biology Knowledgebase



